U.S. DEPARTMENT OF COMMERCE Patent and Trademark Office

SFARCH REQUEST FORM

Requestor's Name:	Serial Number:		
Date:		Ar	t Unit:
Search Topic: Please write a detailed statementerms that may have a special number of the sequence of the sequence.			t matter to be searched. Define any ds, etc., if known. For sequences, levent claim(s).
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PTO-1590 (9-90)



STIC Search Report Biotech-Chem Library

STIC Database Tracking Number: 99905

To: Sarvamangala Devi

Location: CM1-7E12

Art Unit: 1645

Tuesday, July 29, 2003

Case Serial Number: 10/060521

From: Beverly Shears

Location: Biotech-Chem Library

CM1-1E05

Phone: 308-4994

beverly.shears@uspto.gov

Search Notes		
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HarA; enzyme; hygromycin A resistance; bacterial infection; sepsis; septic shock; bacterial endocarditis; prostatitis; NTPase; staphylococcal infection; pneumococcal infection.

B. faecalis hygromycin A resistance protein, harA

20-NOV-2002

(first entry)

ABG32864 standard; Protein; 498

B

Enterococcus faecalis strain OG1X.

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Minimum DB
Maximum DB
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SUMMARIES
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N-PSDB; ABS53679.

2002-676579/73.

Baima ET,

Mueller JP;

30-JAN-2001; 2001US-265034P 29-JAN-2002; 2002BP-0250612

(PFIZ) PFIZER PROD INC.

31-JUL-2002 EP1227324-A2

Novel hygromycin A resistance polypeptide derived from Enterococcus faecalis and Bacillus subtilis useful for screening compounds which binds to the polypeptide, and as medicament for treating drug resistant

Claim 12; Fig 2; 43pp; English.

The invention relates to a recombinant hygromycin A resistance (harA)
CC polypeptide comprising a sequence which is at least 75% identical to
CC the harA protein form Enterococcus faecalis and Bacillus subtilis.
CC Also included are: (1) use of a hygromycin A-resistant strain of
CC Enterococcus faecalis or Bacillus subtilis to determine whether an
CC antibacterial agent is effective in treating organisms which exhibit
CC harA-mediated drug resistance; (2) the nucleic acid encoding the harA
CC polypeptides and (3) harA expression vectors and host cells.
CC HarA protein or nucleic acid is useful in the screening/manufacture of
CC a compound for use as a medicament for the treatment of a drug resistant
CC infection. HarA nucleic acid is useful to identify an organism containing
CC a harA gene. HarA protein is useful to screen for E. faecalis antibodies
CC in blood or serum and samples from animals, or as antigens to raise
CC intibodies, which antibodies are useful as diagnostic reagents. The
CC identified compounds are useful in the treatment of enterococcal
CC infections (e.g. sepsis, septic shock, bacterial endocarditis and
CC prostatitis) as well as staphylococcal and pneumococcal infections.
CC RNA, cDNA and genomic DNA to isolate full-length cDNAs and genomic
CC closes servedity hark polyperit des and to isolate cDNA and genomic

ALIGNMENTS

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B. faecalis hygrom
Enterococcus faeca
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Sequence

498

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Length 498;

harA protein.

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HDAHFMKKI TDKKI VLKS
                                         THRIEOMSMGQRKKVEVAKSLSQSAELYIMDEPLNYLDVFNHQQLEALILSVKPAMLVIB
                                                        TNRIBQMSMGQRKKVBVAKSLSQSABLYIWDBPLNYLDVPNHQQLEALILSVKPAMLVIB
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                                                                                                NPSGDSBGRATLAHQLTISYVRQDYBDNQGTLSBFABKNQLDYTQFLANLRKLGMERAVF
                                                                                                                              SMDYQPTHHKTLLTVEELRLGYEKNWLFAPISPSINAGEIVGITGKNGSGKSSLIQYLLD
                                                                                                                                       SMDYQPTHHKTLLTVBELRLGYEKNMLFAPLSFSINAGEIVGITGKNGSGKSSLIQYLLD
                                                                                                                                                                        EGDKYGNAKEKGSGA I FDYGA I GARAARVMKRSKH I QQRAETQLAEKBKLLKDLEY I DSL
                                                                                                                                                                                   EGDKYGNAKEKGSGAIFDTGAIGARAARVMKRSKHIQQRAETQLAEKEKLLKDLEYIDPL
                                                                                                                                                                                                                  DHILAIBKSQLTLYQGNFSIYBBQKKLRDAFBLAENBKIKKBVNRLKBTARKKABWSMNR
                                                                                                                                                                                                                              DHILAIEKSQLTLYQGNFSIYEBQKKLRDAFELAENEKIKKEVNRLKETARKKAEWSMNR
                                                                                                                                                                                                                                                                                                       498 AA;
                                                                                                                                                                                                                                                                                                                                                                                              Conservative
                                                                                                                                                                                                                                                                                                                                                                                                       99.3%;
                                                                                                                                                                                                                                                                                                                                                                                              <u>ب</u>
                    498
                                                                                                                                                                                                                                                                                                                                                                                                        Score 2522;
Pred. No. 5
                                                                                                                                                                                                                                                                                                                                                                                              Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                     DB 22;
                                                                                                                                                                                                                                                                                                                                                                                                 Indels
                                                                                                                                                                                                                                                                                                                                                                                                                   Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     as drug targets
ints. Cells
                                                                                                                                                                                                                                                                                                                                                                                                                     498;
                                                                                                                                                                                                                                                                                                                                                                                               0,
                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                 360
                                                                                                                                                                                                                                                                                                                                                     60
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                                                                                                                                                                             300
                                                                                                                                                                                                300
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                                                                                                                                                                                                                                          240
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                                                                                                                                                                                                                                                                                                           120
                                                                                                                                                                                                                                                                                                                                120
                                                                                                                                                                                                                                                                                                                                                                           60
                                                                   480
                                                                                                            420
                                              480
                                                                                       420
                                                                                                                                  360
```

480

420

480

420 360 360 300 300

Search Job tim time completed: July me : 99 secs 28, 2003, 15:39:59

481

498

The invention relates to determining whether a candidate nucleotide (ABA82938-ABA82971) or polypeptide (ABA9236-ABB47296) encodes/functions as a multidrug resistance (MDR) efflux pump comprising, searching a database for sequences high identity known MDR efflux pumps and then deleting/mutating an identified region of the DNA in a bacterial cell and determining whether the bacterial cell exhibits increased or decreased sensitivity to an antibacterial agent The identified pumps are useful for

Claim

10; Fig 26; 139pp; English.

```
Result
No.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Title:
Perfect score:
Sequence:
                                                                                                                                                                                                                                                                                                                                                                                                                      Database :
                                                                                                                                                                                                                                                                                                                                                                                                                                                     Post-processing: Minimum Match 75%
Maximum Match 100%
Listing first 1000 summaries
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          OM protein - protein search, using sw model
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Minimum DB seq length: 0
Maximum DB seq length: 2000000000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Scoring table:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Searched:
matches found
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         al number of hits satisfying chosen parameters:
                                                1t Query
O. Score Match Length DB
                                                                                                                                                                               Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.
                                                                                                                                                                                                                                                                      Issued_Patents_AA:*

1: /cgn2_6/ptodata/1/iaa/5A_COMB.pep:*

2: /cgn2_6/ptodata/1/iaa/5B_COMB.pep:*

3: /cgn2_6/ptodata/1/iaa/6A_COMB.pep:*

4: /cgn2_6/ptodata/1/iaa/6B_COMB.pep:*

5: /cgn2_6/ptodata/1/iaa/PCTUS_COMB.pep:*

6: /cgn2_6/ptodata/1/iaa/backfiles1.pep:*
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       328717 seqs, 42310858 residues
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BLOSUM62
Gapop 10.0 , Gapext 0.5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              July 28, 2003, 15:40:06 ; Search time 30 Seconds (without alignments) 702.360 Million cell updates/sec
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        US-10-060-521-2
2540
1 MSKIELKQLSFAYDNQEVLL.....IEHDAHFMKKITDKKIVLKS 498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GenCore version 5.1.6
Copyright (c) 1993 - 2003 Compugen Ltd.
                                                                   Ħ
                                                                                                                                      SUMMARIES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0
                                                                Description
```

Search completed: July 28, 2003, 15:50:53 Job time : 30 secs

```
GenCore version 5.1.6 Copyright (c) 1993 - 2003 Compugen Ltd.
```

OM protein - protein search, using sw model

July 28, 2003, 15:38:06 ; Search time 53 Seconds (without alignments) 1115.894 Million cell updates/sec

Title: Perfect score: US-10-060-521-2 2540

1 MSKIELKQLSFAYDNQEVLL.....IEHDAHFMKKITDKKIVLKS 498

Scoring table:

Searched:

BLOSUM62 Gapop 10.0 , Gapext 0.5

al number of hits satisfying chosen parameters:

0

451899 seqs, 118759770 residues

Minimum DB seq length: 0 Maximum DB seq length: 2000000000

Post-processing: Minimum Match 75%

Maximum Match 100%

Listing first 1000 summaries

Database : Published Applications AA:*

1: /cgn2_6/ptodata/2/pubpaa/US07 PUBCOMB.pep:*

2: /cgn2_6/ptodata/2/pubpaa/US06 NEW PUB.pep:*

3: /cgn2_6/ptodata/2/pubpaa/US06 NEW PUB.pep:*

4: /cgn2_6/ptodata/2/pubpaa/US06 NEW PUB.pep:*

5: /cgn2_6/ptodata/2/pubpaa/US07 NEW PUB.pep:*

6: /cgn2_6/ptodata/2/pubpaa/US08 NEW PUB.pep:*

7: /cgn2_6/ptodata/2/pubpaa/US08 PUBCOMB.pep:*

8: /cgn2_6/ptodata/2/pubpaa/US08 PUBCOMB.pep:*

9: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

10: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

11: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

12: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

13: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

14: /cgn2_6/ptodata/2/pubpaa/US09 PUBCOMB.pep:*

15: /cgn2_6/ptodata/2/pubpaa/US09 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US09 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US10 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US10 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US10 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US09 NEW PUB.pep:*

17: /cgn2_6/ptodata/2/pubpaa/US10 NEW PUB.pep:*

16: /cgn2_6/ptodata/2/pubpaa/US09 NEW PUB.pep:*

17: /cgn2_6/ptodata/2/pubpaa/US09 NEW PUB.pep:*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Result No. Query Score Match Length DB ij Description

No matches found

Search completed: July 28, 2003, 15:48:26 Job time : 53 secs

GenCore version 5.1.6 Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on:

July 28, 2003, 15:36:25; Search time 52 Seconds (without alignments) 921.000 Million cell updates/sec

Title: Perfect score: Sequence:

BLOSUM62 Gapop 10.0 , Gapext 0.5

US-10-060-521-2
2540
1 MSKIELKQLSFAYDNQEVLL.....IEHDAHFMKKITDKKIVLKS 498

Scoring table:

Searched: 283308 seqs, 96168682 residues

al number of hits satisfying chosen parameters:

0

Minimum DB seq length: 0
Maximum DB seq length: 2000000000

Post-processing: Minimum Match 75%

Maximum Match 100%

Listing first 1000 summaries

Database :

PIR 76:*
1: pir1:*
2: pir2:*
3: pir3:*
4: pir4:*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Result No. % Query Score Match Length DB ij

Description

No matches found

ch completed: July 28, 2003, 15:40:58 time : 52 secs

GenCore version 5.1.6 Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

July 28, 2003, 15:35:01; Search time 25 Seconds (without alignments) 936.772 Million cell updates/sec

Run on:

Title: Perfect score: Sequence:

US-10-060-521-2
2540
1 MSKIELKQLSFAYDNQEVLL.....IEHDAHFMKKITDKKIVLKS 498

Scoring table: BLOSUM62
Gapop 10.0 , Gapext 0.5

127863 seqs, 47026705 residues

Searched:

al number of hits satisfying chosen parameters: 0

Minimum DB seg length: 0
Maximum DB seg length: 2000000000

Post-processing: Minimum Match 75%

Maximum Match 100%

Listing first 1000 summaries

Database :

SwissProt_41:*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Result No. Query Score Match Length DB ID

Description

No matches found

Search completed: July 28, 2003, 15:38:21 Job time: 33 secs

```
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```

OM protein - protein search, using sw model

Run on: July 28, 2003, 15:38:26 ; Search time 103 Seconds (without alignments) 1247.672 Million cell updates/sec

US-10-060-521-2 2540

Title: Perfect score:

Sequence: 1 MSKIELKQLSFAYDNQEVLL.....IEHDAHFMKKITDKKIVLKS 498

Scoring table:

BLOSUM62 Gapop 10.0 , Gapext 0.5

Searched: 830525 seqs, 258052604 residues

al number of hits satisfying chosen parameters:

0

Minimum DB seq length: 0 Maximum DB seq length: 200000000

Post-processing: Minimum Match 75%
Maximum Match 100%
Listing first 1000 summaries

Database : SPTREMBL 23:*

1: sp_archea:*

2: sp_bacteria:*

3: sp_fungi:*

sp_invertebrate:*
sp_mammal:*
sp_mhc:* human:*

sp_organelle:*
sp_phage:*
sp_plant:*
sp_plant:*
sp_virus:*
sp_vertebrate:*
sp_writebrate:*
sp_writus:*
sp_rvirus:*
sp_archeap:*

SUMMARIES

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

Result No. Query Score Match Length DB ij Description

No matches found

Search completed: July 28, 2003, 15:50:22 Job time : 110 secs